	<u> </u>
COMPONENTS:	ORIGINAL MEASUREMENTS:
(1) Acetamide. N-[4[[(5-ethyl-1,3,4-thia-	Durel, M. P.; Allinne, M.
diazol-2-yl)amino]sulfonyl]phenyl]-	Bull. Soc. Med. Hop. Paris III
(acetyl sulfaethylthiadiazole);	<u>1941</u> , 251-9.
C <sub>12</sub> H <sub>14</sub> N <sub>4</sub> O <sub>3</sub> S <sub>2</sub> ; [1037-51-0]	
(2) Water; H <sub>2</sub> 0; [7732-18-5]	
VARIABLES:	PREPARED BY:
One temperature: 37°C	R. Piekos
one competators. 37 o	120.103
EXPERIMENTAL VALUES:	
Solubility of acetyl sulfaethylthiadia	zole in water at $37^{\circ}$ C is 0.20 $\circ$ /liter
$(6.1 \times 10^{-4} \text{ mol dm}^{-3}, \text{ compiler}).$	
AUVITTANV	TATABANAMAN
	INFORMATION
METHOD/APPARATUS/PROCEDURE:	SOURCE AND PURITY OF MATERIALS:
A mixt of acetyl sulfaethylthiadiazole and	Source and purity of acetyl sulfaethyl-
water was agitated for 24 hours at 37°C.	thiadiazole was not specified.
	Distilled water was used.
	ESTIMATED ERROR:
	Nothing specified.
	REFERENCES:
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#### ORIGINAL MEASUREMENTS: COMPONENTS: (1) Acetamide, N-[4-[[(5-ethyl-1,3,4-thia-Langecker, H. diazo1-2-v1)amino|sulfonv1|phenv1|-Arch. Exptl. Path. Pharmakol. 1948, (acetyl sulfaethylthiadiazole); 205, 291-301. $C_{12}H_{14}N_4O_3S_2$ ; [1037-51-0] (2) Water: H20; [7732-18-5] VARIABLES: PREPARED BY: R. Piekos рΗ

#### EXPERIMENTAL VALUES:

_ **		Solubility at 37°C			
pH	mg%	10 <sup>4</sup> mol dm <sup>-3</sup> a			
5.2	12	3.7			
6.0	16	4.9			

a Calculated by compiler

# AUXILIARY INFORMATION

# in water was boiled for 1 h in a sealed ampul followed by keeping the ampul at 37°C. Before the assaying, the solute was treated with 2.6N NaOH soln (1) to cleave the acetyl group and the sulfaethylthiadiazole was

An excess of acetyl sulfaethylthiadiazole

METHOD/APPARATUS/PROCEDURE:

detd colorimetrically by the method of Brat-ESTIMATED ERROR: ton and Marshall (2) using a Havemann colorimeter (3), as well as by microanal detd of the solid residue.

# SOURCE AND PURITY OF MATERIALS:

Source and purity of the materials were not specified.

Nothing specified.

# REFERENCES:

- 1. Scudi, J.V. J. Lab. Clin. Med. 1940, 25, 404.
- 2. Bratton, A. G.; Marshall, E.K., Jr. J. Biol. Chem. 1939, 128, 537.
- 3. Havemann, R. Klin. Wochenschr. 1940, p. 503.

- (1) Acetamide, N-[4-[[(5-ethyl-1,3,4-thia-diazol-2-yl)amino]sulfonyl]phenyl](acetyl sulfaethylthiadiazole);
  C12H14N4O3S2; [1037-51-0]
- (2) Phosphoric acid, disodium salt; Na<sub>2</sub>HPO<sub>4</sub>; [7558-94-4]
- (3) Water; H<sub>2</sub>O; [7732-18-5]

# ORIGINAL MEASUREMENTS:

Krüger-Thiemer, E.

Arch. Dermatol. Syphilis <u>1942</u>, 183, 90-116.

#### VARIABLES:

One temperature: ca 20°C; one pH: 8.74

#### PREPARED BY:

R. Piekos

#### EXPERIMENTAL VALUES:

Solubility of acetyl sulfaethylthiadiazole in a 0.705M (10%)  $Na_2HPO_4 \text{ solution of pH 8.74 at room temperature ( about 20°C )}$  is 1.840 g% ( 5.637 x  $10^{-2}$  mol dm<sup>-3</sup> solution, compiler ).

## AUXILIARY INFORMATION

#### METHOD/APPARATUS/PROCEDURE:

Acetyl sulfaethylthiadiazole (0.5 g) was dissolved in 10 cm<sup>3</sup> of the 0.705M (10%) Na<sub>2</sub>HPO<sub>4</sub> soln, shaken for 2 h at room temp (about 20°C), and filtered. The filtrate was treated with equal vol of 2N HCl, and refluxed for 15 min. After proper diln, a 1-cm<sup>3</sup> aliquot was withdrawn, acidified, cooled, and the sulfonamide content was detd colorimetrically (as sulfaethylthiadiazole) by the Marshall method modified by Kimmig (1) using an Authenrieth colorimeter. The pH was detd on an ultraionograph using a glass electrode.

# SOURCE AND PURITY OF MATERIALS:

Acetyl sulfaethylthiadiazole (source not specified) gave no coloration upon diazotization of its satd soln, thus showing absence of sulfaethylthiadiazole. The source and purity of the remaining materials were not specified.

# ESTIMATED ERROR:

Soly: precision ±5% (author).

Temp: not specified.

pH : ±0.05 pH unit (author).

# REFERENCES:

Kimmig, J. Arch. Dermatol. 1938,
 176, 722; Erg. Hyg. 1941, 24,
 398.

- (1) Acetamide, N-[4-[[(5-ethyl-1,3,4-thia-diazol-2-yl)amino]sulfonyl]phenyl]-(acetyl sulfaethylthiadiazole); C<sub>12</sub>H<sub>14</sub>N<sub>4</sub>O<sub>3</sub>S<sub>2</sub>; [1037-51-0]
- (2) Phosphoric acid, monopotassium salt; KH<sub>2</sub>PO<sub>4</sub>; [7778-77-0]
- (3) Water; H<sub>2</sub>0; [7732-18-5]

#### VARIABLES:

One temperature: ca 20°C; one pH: 4.37

#### ORIGINAL MEASUREMENTS:

Krüger-Thiemer, E.

Arch. Dermatol. Syphilis 1942, 183, 90-116.

#### PREPARED BY:

R. Piekos

#### **EXPERIMENTAL VALUES:**

Solubility of acetyl sulfaethylthiadiazole in a 0.735M (10%)  $\rm KH_2PO_4$  solution of pH 4.37 at room temperature ( about  $\rm 20^{\circ}C$  ) is 0.0063 g% ( 1.9  $\times$   $\rm 10^{-4}$  mol dm<sup>-3</sup> solution, compiler ).

#### AUXILIARY INFORMATION

# METHOD/APPARATUS/PROCEDURE:

Acetyl sulfaethylthiadiazole (0.5 g) was dissolved in 10 cm<sup>3</sup> of the 0.735M (10%) KH<sub>2</sub>PO<sub>4</sub> soln, shaken for 2 h at room temp (about 20°C), and filtered. The filtrate was treated with equal vol of 2N HCl and refluxed for 15 min. After proper diln, a 1-cm<sup>3</sup> aliquot was withdrawn, acidified, cooled, and the sulfon-amide content was detd colorimetrically (as sulfaethylthiadiazole) by the Marshall method modified by Kimmig (1) using an Authenrieth colorimeter. The pH was detd on an ultraionograph using a glass electrode.

## SOURCE AND PURITY OF MATERIALS:

Acetyl sulfaethylthiadiazole (source not specified) gave no coloration upon diazotization of its satd soln, thus showing absence of sulfaethylthiadiazole. The source and purity of the remaining materials was not specified.

#### ESTIMATED ERROR:

Soly: precision ±5% (author).

Temp: not specified.

pH : ±0.05 pH unit (author).

# REFERENCES:

1. Kimmig, J. Arch. Dermatol. 1938, 176, 722; Erg. Hyg. 1941, 24, 398.

- (1) Acetamide, N-[4-[[(5-ethyl-1,3,4-thiadiazole-2-yl)amino]sulfonyl]phenyl]-(acetyl sulfaethylthiadiazole);  $c_{12}H_{14}N_4O_3S_2;$ [1037-51-0]
- (2) Phosphoric acid, disodium salt; Na<sub>2</sub>HPO<sub>4</sub>; [7558-94-4]
- (3) Phosphoric acid, monopotassium salt; KH<sub>2</sub>PO<sub>4</sub>; [7778-77-0]
- (4) Water; H<sub>2</sub>0; [7732-18-5]

VARIABLES:

Temperature; pH

#### ORIGINAL MEASUREMENTS:

Krüger-Thiemer, E.

Arch. Dermatol. Syphilis 1942, 183, 90-116.

#### PREPARED BY:

R. Piekos

#### EXPERIMENTAL VALUES:

Composition of 1/15M phosphate

buffer colutions

		Solubility
Da	*	(an 200c)

buffer solutions	- pH Room	Room t	emp (ca 20°C)	37°C			
Na <sub>2</sub> HPO <sub>4</sub>	кн <sub>2</sub> РО <sub>4</sub>	%Content	pii	g%	10 <sup>3</sup> mol dm <sup>-3</sup> solution a	g%	10 <sup>3</sup> mol dm <sup>-3</sup> solution <sup>a</sup>
1.0	99.0	0.91	4.944	0.0128	0.392	-	-
10.0	90.0	0.91	5.906	0.0530	1.600	0.112	3.43
61.1	38.9	0.93	7.005	0.3910	12.0	0.750	22.98
9.5	0.5	0.733 <sup>b</sup>	7.51	1.1100	34.01	-	-
94.7	5.3	0.95	8.018	0.8790	26.9	-	-

a Calculated by compiler

# AUXILIARY INFORMATION

# METHOD /APPARATUS / PROCEDURE:

Acetyl sulfaethylthiadiazole (0.5 g) was dis Acetyl sulfaethylthiadiazole (source not solved in 10 cm3 of a buffer soln, shaken for specified) gave no coloration upon diazo-2 h at  $20^{\circ}$ C (or left for 48 h at  $37^{\circ}$ C), and filtered at respective temp. The filtrate was treated with equal vol of 2N HCl and refluxed for 15 min. After proper diln, a 1-cm<sup>3</sup> aliquot was withdrawn, acidified, cooled, and the sulfonamide content was detd colorimetrically (as sulfaethylthiadiazole) by the Marshall method modified by Kimmig (1) using an Authenrieth colorimeter. The pH was detd on an ultraionograph using a glass electrode

# SOURCE AND PURITY OF MATERIALS:

tization of its satd soln, thus showing absence of sulfaethylthiadiazole. The source and purity of the remaining materials were not specified.

## ESTIMATED ERROR:

Soly: precision ±5% (author).

Temp: not specified.

pH : ±0.05 pH unit (author).

#### REFERENCES:

1. Kimmig, J. Arch. Dermatol. 1938, 176, 722; Erg. Hyg. 1941, 24, 398.

b Molar content; 10% buffer solution

- (1) Acetamide, N-[4-[(5-ethyl-1,3,4-thiadiazol-2-yl)amino]sulfonyl]phenyl]-(acetyl sulfaethylthiadiazole);  $C_{12}H_{14}N_4O_3S_2$ ; [1037-51-0]
- (2) Phosphoric acid, disodium salt; Na<sub>2</sub>HPO<sub>4</sub>; [7558-94-4]
- (3) Phosphoric acid, monopotassium salt; KH<sub>2</sub>PO<sub>4</sub>; [7778-77-0]
- (4) Water; H<sub>2</sub>0; [7732-18-5]

VARIABLES:

pН

#### ORIGINAL MEASUREMENTS:

Bandelin, F. J.; Malesh, W. J. Am. Pharm. Assoc., Sci. Ed. 1959, 48, 177-81.

PREPARED BY:

R. Piekos

#### EXPERIMENTAL VALUES:

Solubility of acetyl sulfaethylthiadiazole in buffers of varying mixtures of  $Na_2HPO_4 \cdot 7H_2O$  (71.6 g/l distilled water; 0.27 mol dm<sup>-3</sup>, compiler) and  $KH_2PO_4$ (36.3 g/l distilled water; 0.27 mol  $dm^{-3}$ , compiler) at 37°C.

Solubility (based on sulfaethylthiadiazole)

		·,
Equilibrium pH	mg/100 m1	$10^2$ mol dm <sup>-3</sup> a
4.5	140	0.492
4.6	162	0.570
5.2	212	0.745
5.6	300	1.055
6.2	510	1.794
6.6	740	2.602
6.8	1175	4.132

<sup>&</sup>lt;sup>a</sup> Calculated by compiler

#### AUXILIARY INFORMATION

# METHOD/APPARATUS/PROCEDURE:

Solns were prepd by adding an excess of ace- | Neither source nor purity of the reagents tyl sulfaethylthiadiazole to 10 ml of buffer were specified. Distilled water was used. soln at each pH level in 18 x 150-mm test tubes, stoppering the tubes, and placing them in water bath at 37°C with gentle agitation for 24 h. The solute was then hydrolyzed with 5%  $H_2SO_4$  for 1 h to liberate the free sulfonamide. One-ml aliquot of the hydrolyzate was accurately pipetted into a volumetric flask for diln and analysis. The sulfonamide was assayed colorimetrically by the method of Bratton and Marshall as described in detail by Biamonte and Schneller (1). A standard curve was prepd using accurately prepd standard solutions.

## SOURCE AND PURITY OF MATERIALS:

# ESTIMATED ERROR:

Soly: av values of ducplicate runs are reported (authors).

Temp and pH: not specified.

# REFERENCES:

1. Biamonte, A. R.; Schneller, G. E. J. Am. Pharm. Assoc., Sci. Ed. 1952, 41, 341.

- (1) Acetamide, N-[4-[[(5-ethyl-1,3,4-thia-diazol-2-yl]amino]sulfonyl]phenyl](acetyl sulfaethylthiadiazole);
  C12H14N403S2; [1037-51-0]
- (2) Phosphoric acid, disodium salt; Na<sub>2</sub>HPO<sub>4</sub>; [7558-94-4]
- (3) Phosphoric acid, monopotassium salt; KH<sub>2</sub>PO<sub>4</sub>; [7778-77-0]
- (4) Water; H<sub>2</sub>0; [7732-18-5]

VARIABLES:

pН

# ORIGINAL MEASUREMENTS:

Hekster, Ch. A.; Vree, T. B.

Antibiotics Chemother. 1982, 31,
22-118.

PREPARED BY:

R. Piekos

#### EXPERIMENTAL VALUES:

	Solubility at 25°C			
рН	mg/l	10 <sup>3</sup> mol dm <sup>-3</sup> a		
5.5	392	1.20		
7.5 <sup>b</sup>	7,850	24.05		

<sup>&</sup>lt;sup>a</sup>Calculated by compiler

in the article

#### AUXILIARY INFORMATION

# METHOD/APPARATUS/PROCEDURE:

The earlier developed method (1) was used (personal communication). Satd solns of acetyl sulfaethylthiadiazole were prepd in phosphate buffers of pH 5.5 and 7.5 at 25°C. The concn of the solute was measured by means of a Spectra Physics 3500B high-performance liquid chromatograph equipped with a Model 748 column oven and a Pye-Unicam LC-UV spectrophotometric detector.

#### SOURCE AND PURITY OF MATERIALS:

Neither source nor the purity of the materials was not specified.

#### ESTIMATED ERROR:

Soly: the detection limit of the solute by HPLC was 0.5 mg/l (authors).

The errors in temp and pH were not specified.

# REFERENCES:

Hekster, Y. A.; Vree, T. B.;
 Damsma, J. E.; Friesen, W. T.
 J. Antimicrob. Chemother. 1981, 8,
 133.

bErroneous pH value of 7.0 is given